

WHAT IS CLAIMED IS:

1. A method of fabricating an array of polynucleotides on a substrate, comprising:
 - (a) operating a polynucleotide deposition system to deposit an array of polynucleotide containing fluid droplets on the substrate to provide, when dry, a target pattern of polynucleotide containing spots;
 - (b) allowing a sufficient time to pass such that droplets deposited by the system will have dried to yield an actual pattern of dried spots; and
 - (c) observing the actual pattern; and
 - (d) comparing the actual pattern with the target pattern of polynucleotide containing spots.
2. A method according to claim 1 wherein the actual pattern is observed by capturing an image of the actual pattern.
3. A method according to claim 1 wherein the system deposits fluid droplets at least some of which contain respective different polynucleotides.
4. A method according to claim 1 wherein the one or more polynucleotide containing fluids each comprise a solution of a polynucleotide and a sufficient amount of salt to enhance polynucleotide imaging.
5. A method according to claim 1 wherein the polynucleotides are DNA of at least six nucleotides in length.
6. A method according to claim 5 wherein the DNA is cDNA.
7. A method according to claim 5 wherein the DNA is single stranded.

8. A method according to claim 1 wherein the comparison includes a comparison of dried actual spot locations or dimensions with target locations or dimensions.
9. A method according to claim 1 wherein the comparison includes a comparison of the presence or absence of dried spots at target locations.
10. A method according to claim 1 wherein the image capture comprises imaging a light scattering characteristic of dried spots.
11. A method according to claim 1 wherein the image capture comprises imaging a fluorescence characteristic of dried spots.
12. A method according to claim 1 additionally comprising generating a signal indicative of the result of the comparison.
13. A method according to claim 1 wherein the deposition system is operated to fabricate multiple polynucleotide arrays on a same substrate.
14. A method according to claim 1 additionally comprising:
when the results of one or more comparisons for an array exceed a predetermined tolerance, storing an error indication in association with that array.
15. A method according to claim 14 wherein the deposition system is operated to fabricate multiple polynucleotide arrays, the method additionally comprising rejecting the array associated with a stored error indication.
16. A method according to claim 14 wherein the error indication is stored in a memory, the method additionally comprising writing on a medium an identification for the array associated with an error indication; physically

associating that medium with the array; and storing the identification in the memory.

17. A method according to claim 16 wherein the error indication includes an indication of the magnitude of the error.

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18. A method of fabricating an array of polynucleotides on a substrate, comprising:

- (a) operating a polynucleotide deposition system to deposit an array of polynucleotide containing fluid droplets on the substrate to provide a target pattern of polynucleotide containing spots which may differ from the actual pattern deposited;
- (b) capturing an image of the actual pattern;
- (d) comparing the actual pattern with the target pattern of polynucleotide containing spots;
- (e) when the results of one or more comparisons for an array exceed a predetermined tolerance, generating an error indication associated with the array;
- (f) writing the error indication or an identification of the error indication, on a medium; physically associating the medium with the array; and, when an identification of the error indication is written on the medium, storing the identification in the memory in association with the error indication.

19. A method of identifying an error in an array of polynucleotides on a substrate, which array is associated with a medium carrying an identification of an error indication, the method comprising communicating the identification to a remote server on which is stored the identification and the error indication, receiving in response the error indication from the remote server, exposing the array to a biological sample, determining an observed binding pattern of the exposed array, and modifying the determination or altering the results of the determination based on the received error indication.

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~~20~~. A method according to claim 1 wherein the polynucleotide deposition system automatically fabricates multiple polynucleotide arrays, the method additionally comprising, when the results of one or more comparisons for an array exceed a predetermined tolerance indicating an error condition, automatically halting further operation of the deposition system and generating a visible or audible operator alert.

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~~21~~. A method according to claim 1 wherein the polynucleotide deposition system includes a fluid dispensing head with multiple drop dispensers, the method additionally comprising:

when the results of one or more comparisons for an array exceed a predetermined tolerance, generating an error indication;

when multiple error conditions are generated, evaluating if a same drop dispenser is responsible;

and wherein, when the evaluation indicates the same drop dispenser is responsible, generating a visible or audible operator alert which includes an indication of the responsible drop dispenser.

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~~22~~. A method according to claim ¹⁹~~21~~ wherein the indication of the responsible drop dispenser comprises an indication of potential error in a polynucleotide containing fluid which has been preselected to be dispensed by that drop dispenser.

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~~23~~. A method according to claim 1 wherein the polynucleotide deposition system includes a fluid dispensing head with multiple drop dispensers and a control processor, the method additionally comprising:

the control processor loading the dispensers in a pattern in which at least some of the dispensers are loaded with the same fluid;

when the results of one or more comparisons for an array exceed a predetermined tolerance, generating an error indication;

when multiple error indications are generated, the control processor comparing a pattern of error indications with the loading pattern of the dispensers to evaluate whether one or more drop dispensers or an error in a polynucleotide containing fluid is responsible for the error indications.

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24. A method according to claim 13 wherein the polynucleotide deposition system includes a fluid dispensing head with multiple drop dispensers, and each array is to be deposited using an initial pattern of drop dispensing from the multiple dispensers, the method additionally comprising:

when the results of one or more comparisons for an array exceed a predetermined tolerance, generating an error indication;

when multiple error indications are generated, evaluating if a same drop dispenser may be responsible; and

when the evaluation indicates the same drop dispenser may be responsible, altering the initial pattern such that the same drop dispenser is not used.

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25. An apparatus for fabricating an array of polynucleotides on a substrate, comprising:

- (a) a polynucleotide deposition system to deposit an array of polynucleotide containing fluid droplets on the substrate to provide, when dry, a target pattern of polynucleotide containing spots;
- (c) an imaging system to capture an image of an actual pattern of dried spots resulting from drying of droplets deposited on the substrate;
- (d) a processor to control the deposition system to deposit the array of droplets and which, after a predetermined time has elapsed for drying of the droplets to yield the actual pattern, causes the imaging system to capture the actual pattern and compares actual pattern with the target pattern of polynucleotide containing spots.

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26. An apparatus according to claim ²⁸25 wherein the deposition system deposits droplets of no greater than 1000 pL.

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~~25.~~ An apparatus according to claim ~~25~~²⁸ wherein the polynucleotide deposition system includes a fluid dispensing head with multiple drop dispensers, and wherein the processor:

when multiple error conditions are generated, evaluates if a same drop dispenser may be responsible.

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36. An apparatus according to claim 36 additionally comprising an audio or visual output device, and wherein when the evaluation indicates a same drop dispenser may be responsible the processor generates an operator alert on the output device which includes an indication of the responsible drop dispenser.

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evaluates if a same drop dispenser or an error in a polynucleotide containing fluid may be responsible by comparing the stored dispensing pattern with a pattern of error indications.

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